

Claims

1. A method of coating the internal surface of a device with a polymer, the process comprising the steps of:

- 5 (i) introducing into the device a solution of one or more monomers in a suitable solvent;
(ii) introducing a flow of an inert gas through the device; and
(iii) initiating polymerisation of the monomer solution.

2. A method as claimed in claim 1 wherein the device is a microfabricated device or a
10 reaction vessel with an internal diameter of less than about 2mm

3. A method as claimed in claim 1 or claim 2, wherein the inert gas is nitrogen or argon.

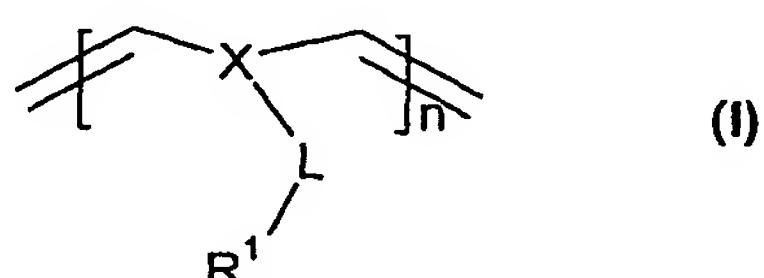
15 4. A method as claimed in any one of claims 1 to 3, wherein the device is a microfabricated device or a loop from 1 to 100 cm in length.

5. A method as claimed in any one of claims 1 to 4, wherein the device is adapted to carry out a solid-phase radiochemical process.

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6. A method as claimed in any one of claims 1 to 5, wherein the one or more monomers can be polymerised by ring opening metathesis polymerisation (ROMP) and the solution also includes a ruthenium carbene catalyst and a cross-linker.

25 7. A method as claimed in any one of claims 1 to 6, wherein polymerisation of the one or more monomers leads to a ROMP polymer of Formula (I):



wherein:

30 X is either a C₄₋₆ cycloalkyl or C₄₋₆ heterocyclyl moiety;

L is a C₁ to C₂₀ linker group comprising one or more alkyl, alkenyl, alkynyl, C₄₋₁₀ cycloalkyl, C₄₋₁₀ heterocyclyl, C₄₋₁₀ aryl, C₄₋₁₀ heteroaryl, ether, PEG, sulphide, amide, sulphamide or a combination thereof; any of which may be substituted with one or more groups R²

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R¹ is hydrogen, C₁₋₂₀ alkyl, C₂₋₂₀ alkenyl, C₂₋₂₀ alkynyl, C₄₋₁₂ cycloalkyl, C₄₋₁₂ heterocyclyl, aryl, heteroaryl, C(O)R³, C₁₋₂₀ alkyl-C(O)R³, C₂₋₂₀ alkenyl-C(O)R³, C₂₋₂₀ alkynyl-C(O)R³, nitro, isocyanate, C₁₋₁₀ alkyl-C(O)-C(R⁴)₂-C(O)-C₁₋₁₀ alkyl, aminoxy, nitrile, phosphorus chloride, succinimide, sulphonyl chloride, halogen, tosylate, mesylate, triflate, nonaflate, 10 silane, OR⁴, SR⁴, N(R⁴)₂, N^{+(R⁴)₃, quaternary phosphorous, C₁₋₂₀ alkyl-R⁵, C₂₋₂₀ alkenyl-R⁵ or C₂₋₂₀ alkynyl-R⁵ or a group comprising an enzyme or a catalyst.}

R² is C(O)R³, C₁₋₂₀ alkyl-C(O)R³, C₂₋₂₀ alkenyl-C(O)R³, C₂₋₂₀ alkynyl-C(O)R³, nitro, isocyanate, C₁₋₁₀ alkyl-C(O)-C(R⁴)₂-C(O)-C₁₋₁₀ alkyl, aminoxy, nitrile, phosphorus chloride, 15 succinimide, sulphonyl chloride, halogen, tosylate, mesylate, triflate, nonaflate, silane, OR⁴, SR⁴, N(R⁴)₂, N^{+(R⁴)₃, quaternary phosphorous, C₁₋₂₀ alkyl-R⁵, C₂₋₂₀ alkenyl-R⁵ or C₂₋₂₀ alkynyl-R⁵.}

R³ is H, OH, C₁₋₂₀ alkyl, OC₁₋₂₀ alkyl, N(R⁴)₂, N^{+(R⁴)₃;}

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each R⁴ is independently H or C₁₋₁₀ alkyl;

R⁵ is OR⁴, SR⁴, N(R⁴)₂, N^{+(R⁴)₃, C₄₋₁₀ cycloalkyl, C₄₋₁₀ heterocyclyl, aryl or heteroaryl.}

25 8. A process as claimed in claim 7, wherein, in the ROMP polymer of Formula (I):

R¹ is halogen, OH, SH, C₁₋₂₀ alkyl, C₄₋₁₂ aryl, C₁₋₂₀ alkyl-R⁵, C₁₋₂₀ alkyl-C(O)R³, N(R⁴)₂, N^{+(R⁴)₃ or a group comprising an enzyme or a catalyst.}

where R³ is OH, R⁴ is as defined for general formula (I) and R⁵ is N(R⁴)₂, N^{+(R⁴)₃, aryl or heteroaryl;}

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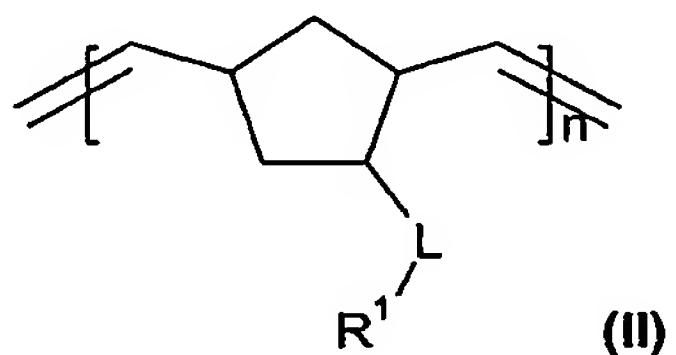
9. A process as claimed in claim 8, wherein, in the ROMP polymer of Formula (I) wherein R¹ is C₁₋₂₀ alkyl; -N=C=O, -SH or N^{+(R⁴)₃, particularly with bound ¹⁸F-fluoride ion}

or comprises an enzyme or a catalyst; and R⁴ is as defined in general formula (I).

10. A process as claimed in any one of claims 7 to 9, wherein the polymer of Formula (I) contains more than one R¹ group.

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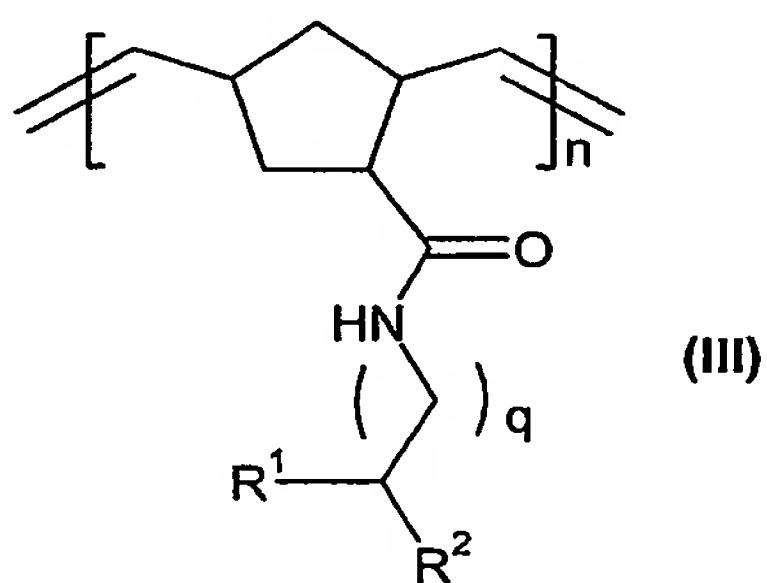
11. A process as claimed in any one of claims 1 to 10 wherein polymerisation of the one or more monomers leads to a ROMP polymer of Formula (II):



10 wherein:

-L-, R¹ and n are as defined above for Formula (I).

12. A process as claimed in any one of claims 1 to 11 wherein polymerisation of the one or more monomers leads to a ROMP polymer of Formula (III):



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wherein:

R¹ and n are as defined above for Formula (I);

R² is an optional group as defined above for -L- of Formula (I); and,

q = 1-4.

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13. A process as claimed in claim 12, wherein, in the ROMP polymer of Formula (III), R¹ is trialkylammonium, R² is absent, q = 3 and n = number of polymer units.

14. A process as claimed in any one of claims 1 to 13, wherein each monomer is

present in the starting solution in a concentration of from about 0.1 to 5M.

15. A process as claimed in any one of claims 1 to 14 wherein, in the monomer solution, the solvent is a polar aprotic solvent.

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16. A process as claimed in any one of claims 1 to 15 wherein polymerisation is initiated by heating.

17. A process as claimed in any one of claims 1 to 15 wherein polymerisation occurs
10 spontaneously.

18. A process as claimed in any one of claims 1 to 17, wherein the device is a microfabricated device and, the process of the invention comprises the initial step of creating a defined network of channels within the device.

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19. A device comprising a microfabricated device or a reaction vessel with an internal diameter of less than about 2mm, wherein the internal surface is coated with a polymer substrate for a solid phase physical or chemical process.

20 20. A device as claimed in claim 19 adapted for carrying out a solid phase radiochemical process.

21. A device as claimed in claim 19 or claim 20, wherein the internal surface is coated . . .
with a ROMP polymer.

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22. A device as claimed in any one of claims 19 to 21, wherein the internal surface is coated with a polymer as defined in any one of claims 7 to 13.

23. An automated synthesis system comprising two or more devices as claimed in any
30 one of claims 19 to 22 which are fluidly interconnected

24. A method for recovering of ^{18}F -fluoride ion from ^{18}O -enriched water containing ^{18}F -fluoride ion, the process comprising passing the ^{18}O -enriched water containing ^{18}F -fluoride

ion through a device as claimed in any one of claims 19 to 22 or a system as defined in claim 23, in which the polymer coating comprises a ROMP polymer of general formula (III) in which R¹ is tri(C₁₋₆ alkyl)ammonium, with a non-nucleophilic counter-ion, R² is absent and q is 3.

- 5 25. A method as claimed in claim 24 which is a step in the synthesis of an ¹⁸F-labelled radiotracer.
26. A method for the synthesis of an ¹⁸F-labelled radiotracer, the method comprising:
- (i) recovering of ¹⁸F-fluoride ion from ¹⁸O-enriched water containing ¹⁸F-fluoride ion passing the ¹⁸O-enriched water containing ¹⁸F-fluoride ion through a device as claimed in
10 any one of claims 19 to 22 or a device as claimed in claim 23 in which the polymer coating comprises a ROMP polymer of general formula (III) in which R¹ is tri(C₁₋₆ alkyl)ammonium, with a non-nucleophilic counter-ion, R² is absent and q is 3; and
- (ii) introducing into the device an unlabelled precursor compound of the ¹⁸F-labelled radiotracer such that ¹⁸F becomes incorporated into the precursor compound *via*
15 nucleophilic substitution to form the ¹⁸F-labelled radiotracer.
27. A method as claimed in claim 26, wherein the ¹⁸F-labelled radiotracer is:
- 2-[¹⁸F]fluorodeoxyglucose (2-[¹⁸F]-FDG);
- L-6-[¹⁸F]fluoro-DOPA;
- 3'-deoxy-3'-fluorothymidine (FLT);
- 20 2-(1,1-dicyanopropen-2-yl)-6-(2-[¹⁸F]fluoroethyl)-methylamino)-naphthalene ([¹⁸F]FDDNP);
- 5[¹⁸F]fluorouracil; 5[¹⁸F]fluorocytosine; or
- [¹⁸F]-1-amino-3-fluorocyclobutane-1-carboxylic acid ([¹⁸F]-FACBC).